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Combined Diet and Physical Activity Promotion Programs to Prevent Type 2 Diabetes Among People at Increased Risk: A Systematic Review for the Community Preventive Services Task Force

Ethan M. Balk, MD, MPH^{1,2}, Amy Earley, BS³, Gowri Raman, MD, MS², Esther A. Avendano, BA², Anastassios G. Pittas, MD, MS⁴, and Patrick L. Remington, MD, MPH⁵

¹Center for Evidence Based Medicine, Brown University School of Public Health, Providence RI

²Center for Clinical Evidence Synthesis, Institute of Clinical Research and Health Policy Study, Tufts Medical Center, Boston MA

³Mapi Group, Boston MA

⁴Division of Endocrinology Diabetes and Metabolism, Tufts Medical Center, Boston MA

⁵School of Medicine and Public Health, University of Wisconsin, Madison WI

Abstract

Background—Trials have demonstrated the efficacy of rigorous diet and physical activity promotion (D&PA) programs for adults at increased risk for type 2 diabetes to reduce diabetes incidence and improve measures of glycemia.

Purpose—To evaluate D&PA programs for individuals at increased risk for type 2 diabetes primarily to lower diabetes risk, lower body weight, and improve glycemia.

Data Sources—MEDLINE, Cochrane Central Register of Controlled Trials, CAB Abstracts, Global Health, and Ovid HealthStar from 1991 through 27 February 2015, with no language restriction.

Corresponding Author: Ethan Balk, MD MPH, Center for Evidence-Based Medicine, Box G-S121-8, Brown University School of Public Health, Providence, RI 02912, USA, ethan_balk@brown.edu.
Current addresses:

Ethan M. Balk, MD, MPH, Brown University School of Public Health, Box G-S121-8, Providence, RI 02912

Amy Earley, BS, Mapi USA, 180 Canal Street, Suite 503, Boston MA 02114, aearley@mapigroup.com

Gowri Raman, MD, MS, Center for Clinical Evidence Synthesis, Tufts Medical Center, 800 Washington Street, Box 063, Boston, MA 02111, graman@tuftsmedicalcenter.org

Esther A. Avendano, BA, Center for Clinical Evidence Synthesis, Tufts Medical Center, 800 Washington Street, Box 063, Boston, MA 02111, eavendano@tuftsmedicalcenter.org

Anastassios G. Pittas, MD, MS, Division of Endocrinology, Diabetes and Metabolism, Tufts Medical Center, 800 Washington Street, Box 268, Boston, MA 02111, apittas@tuftsmedicalcenter.org

Patrick L. Remington MD, MPH, Health Science Learning Center, Rm 4263, 750 Highland Avenue, Madison, WI 53705, plreming@wisc.edu

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Study Selection—8 researchers screened articles for single group or comparative studies of combined D&PA programs with at least 2 sessions of at least 3 month duration in participants at increased risk for type 2 diabetes.

Data Extraction—7 researchers extracted data—on study design, participant, intervention, outcome descriptions, and results—and assessed study quality.

Data Synthesis—53 studies (30 D&PA vs. control, 13 more vs. less intensive, 13 in single programs) evaluated 66 programs. Compared with usual care, D&PA reduced type 2 diabetes incidence (RR = 0.59; 95% CI 0.51, 0.66; 16 studies), lowered body weight (net change = -2.2%; 95% CI -2.9, -1.4; 24 studies) and fasting blood glucose (net change = -0.12 mmol/L; 95% CI -0.20, -0.05; 17 studies), and improved other cardiometabolic risk factors. There was limited evidence for clinical events. More intensive programs were more effective.

Limitations—The wide variation in D&PA programs limited identification of features most relevant to effectiveness. Evidence on clinical outcomes and in children was sparse.

Conclusions—Combined D&PA promotion programs are effective to decrease diabetes incidence and improve cardiometabolic risk factors for patients at increased risk. More intensive programs are more effective.

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Keywords

pre-diabetes; type 2 diabetes; diet; exercise; primary prevention

Diabetes is a large and growing medical problem and the costs to society are high and escalating. According to the latest figures from the Centers for Disease Control and Prevention (CDC), 29.1 million people or 9.3% of the U.S. population have diabetes and 1.7 million new cases are diagnosed annually (1). Worldwide, it is estimated that 387 million adults are living with diabetes and this number is projected to increase to 592 million by 2035 (2). Prevalence of diabetes and related costs are expected to more than double in the next quarter century (3), as more than 86 million Americans (37% of the adult population) are at risk of developing diabetes (1). Effective prevention strategies are, therefore, critically important to slow the diabetes tide and its associated burden.

Nearly 9 out of 10 new cases of diabetes are due to type 2 diabetes, whose natural history is characterized by a gradual rise in glycemia. Identifying those at increased risk can allow the implementation of interventions to lower the risk of progressing to clinical diabetes. The American Diabetes Association has defined pre-diabetes as a high-risk category, based on a level of glycemia that does not meet criteria for diabetes but is too high to be considered normal (4). People with pre-diabetes progress to type 2 diabetes at a rate of about 5–10% per year without any intervention (5). Three large clinical trials from the US (6), Finland (7), and China (8) have shown that the main components of diabetes prevention in adults are weight loss and increased physical activity. In these trials, among people at risk for type 2 diabetes, rigorous application of combined diet and physical activity (D&PA) promotion programs, with the goals of weight loss and increased physical activity were successful at

reducing risk of diabetes by 50–60% during the active intervention period (3 to 6 years). Although attenuated, the effect of the intervention can persist long-term (9–11). The results of these trials are well known; however, wide-scale implementation in clinical and community-based settings has recently begun, but requires further progress (12).

Combined D&PA promotion programs aim to prevent type 2 diabetes among people who are at increased risk for the disease. These programs actively encourage people to improve their diet and increase their physical activity using trained providers in various settings who work with clients for at least 3 months, providing some combination of counseling, coaching, and extended support in multiple sessions related to diet and physical activity, delivered in-person, or by other methods. Programs may also include numerous other features, including specialized counselors, a range of number and frequency of sessions, different session types, and different diet, weight loss, or exercise goals.

The purpose of this review was to assess the effectiveness of D&PA promotion programs implemented in a wide range of clinical or community settings to reduce risk of new-onset diabetes among adults and children at risk for type 2 diabetes. The Community Preventive Services Task Force (Task Force, www.thecommunityguide.org) used this review to update its guidance in diabetes prevention and to identify current gaps in the evidence to inform future research. Potential effect modifiers such as intensity and specificity of the programs, settings and implementers were evaluated. Furthermore, the potential benefit of the diabetes prevention programs extending to other cardiometabolic risk factors, such as overweight, high cholesterol and high blood pressure, was also assessed.

Methods

The review was conducted in accordance with the methodology of the Task Force (13;14) and the highest standards for conducting systematic reviews (15;16). We convened a panel of domain experts and stakeholders (Coordination Team), which together with our Community Guide Technical Monitor and Task Force members provided input regarding the protocol, feedback on the findings, conclusions, and evidence gaps.

Data Sources

We searched MEDLINE, the Cochrane Central Register of Controlled Trials, CAB Abstracts, Global Health, and Ovid HealthStar from 1991 through 27 February 2015 with no language restrictions. Supplemental Table 1 shows the search strategy. We also screened through reference lists of related systematic and narrative reviews, and suggestions from the expert panel.

Study Selection

We included randomized controlled trials and prospective non-randomized comparative studies with at least 30 participants per group, and prospective single group intervention studies with at least 100 participants. The population of interest was focused on adults or children at increased risk for type 2 diabetes (i.e., with pre-diabetes) as determined by glycemia measures or diabetes risk assessment tools. We included studies of participants with metabolic syndrome (who are at increased risk of both diabetes and cardiovascular

disease) or whose participants were chosen because they were at risk for *either* type 2 diabetes or cardiovascular disease. However, we excluded studies of participants with established type 2 diabetes or whose *only* risk factor was obesity or increased risk for cardiovascular disease (without explicit inclusion of participants with pre diabetes). The implied or explicit intent of the D&PA programs had to be to prevent diabetes and had to include at least 2 contact sessions (in-person or virtually) over a minimum period of 3 months. Programs had to include both dietary and increased physical activity components and could be conducted in any outpatient setting. We allowed any type of advice to improve diet and increase physical activity (except single food or supplement dietary changes such as adding fish oil). We excluded interventions that included antidiabetic medications. The comparative studies had to include a usual care arm (no active D&PA program) or a lower-intensity D&PA program (e.g., with fewer contact sessions or a more liberal diet). We required at least 6 month follow-up for any of the following outcomes: incident diabetes, reversion to normoglycemia, body weight, glycemia measures (fasting glucose [FG], 2 hour glucose after a 75-gram oral glucose tolerance test [2hG], hemoglobin A1c [HbA1c]), all-cause death, diabetes-related clinical outcomes (e.g., cardiovascular events, end-stage renal disease, nephropathy, amputation, retinopathy, neuropathy, skin ulcers, periodontitis), blood pressure (BP), and lipids (total cholesterol, low and high density lipoprotein cholesterol [LDL, HDL], and triglycerides).

Data Extraction and Quality Assessment

We screened titles and abstracts using *Abstrackr* (17). Eight researchers double-screened the abstracts after iterative training of all reviewers on the same batches of abstracts. Discordant decisions and queries were resolved at group meetings. Full-text articles were retrieved for all potentially relevant abstracts and rescreened by the same researchers.

Each study was extracted by one of seven experienced methodologists and confirmed by a senior methodologist; the same methodologists assessed study quality. Data extraction was conducted in SRDR (18), and included elements for study design including eligibility criteria, population characteristics, detailed descriptions of the D&PA programs and comparison interventions, outcomes, and results. We assessed each study's quality based on 12 Community Guide quality of execution questions (Supplemental Table 2, footnotes) (14;19). Per Community Guide protocol, we excluded studies with "limited quality of execution," defined as having at least five major limitations.

Data Synthesis and Analysis

All extracted data were tabulated into Summary Evidence Tables (available in the Supporting Materials at the URL listed at the beginning of the Results section). Because only two studies were conducted in children, we report these separately and do not include them in the meta-analyses. For outcomes with data from at least three comparative studies of D&PA vs. usual care, we performed profile likelihood random effects model meta-analysis of risk ratio (RR) or net change (20). For non-randomized studies, we preferentially used adjusted analysis results. Meta-analyses were conducted with the *metaan* package in Stata 13.1 (StataCorp, College Station, TX). For the overall meta-analyses of incident diabetes and reversion to normoglycemia, we used the longest reported duration of follow-up data.

For continuous outcomes, we used the data closest to 1 year of follow-up, data restricted to <2 years of follow-up, and longest followup. We evaluated differences in effect (for incident diabetes and weight only) based on any direct comparisons of different D&PA programs within studies, any reported within-study subgroup analyses, and across-study random effects model metaregression across all D&PA programs based on predetermined study setting and D&PA program features. Incident diabetes and weight change were chosen for metaregression based on their relative importance in determining the effectiveness of D&PA programs. Metaregressions were conducted with the metareg package in Stata and were considered to be potentially significant if $P < 0.10$. For each outcome with at least 10 studies, the possibility of publication bias was examined with funnel plots and the Harbord test for diabetes incidence and the Egger test for continuous outcomes using the metabias and metafunnel packages in Stata (21).

Role of the Funding Source

One member of the Coordination Team and our Technical Monitor are employed by the CDC; none of the Task Force members are. The Coordination Team, Technical Monitor, and members of the Community Preventive Services Task Force participated in formulating the study questions and developing the protocol but did not participate in the literature search, determination of study eligibility criteria, data analysis or interpretation. The Coordination Team, Technical Monitor, and CDC personnel were provided with an opportunity to provide feedback on the manuscript and the decision to submit the manuscript for publication, but the research team retained final determination of the content and decision to publish.

Results

Supplemental Figure 1 summarizes the search yield. Of 11,317 citations (plus articles from other existing systematic reviews and domain experts), 53 studies described 66 D&PA programs in 104 articles (6–11;22–119). One additional study was excluded for limited quality of execution (with six major limitations) (120). The included studies described 26 randomized and 4 nonrandomized comparisons of D&PA programs versus usual care, 12 randomized and 1 nonrandomized comparisons of two or more D&PA programs (3 of which also had usual care arms), and 13 single group studies of D&PA programs. Thirty-three studies were of good quality (0 to 1 limitation) and 20 were of fair quality (2 to 4 limitations) (Supplemental Table 2). The most common limitations were poor descriptions of the study populations or intervention programs; problems with data measurement or interpretation; and high dropout rates. While half the studies ($n=27$) analyzed all enrolled participants, nine had more than 20% drop-out (or loss-to-follow-up) rates.

The characteristics of the D&PA programs are summarized in Table 1 (with details in Supplemental Tables 3–5). All but five programs (in four studies) ran for at least 6 months. Programs offered a wide range of number of contact sessions (0 [virtual contacts only] to 72, median 15) and most programs included both a core period (with frequent contact sessions) and a maintenance period (when participants were contacted less frequently). With the exception of seven programs that were delivered entirely over the internet, by video,

telephone, or email, programs used in-person individual or group sessions, or both, on diet or exercise, or both. Sessions were led by different combinations of trained diet counselors including dietitians, nutritionists, or others; trained exercise counselors including physical trainers or others; nurses; physicians or psychologists; or trained laypeople. Many programs included specific weight loss, diet, or physical activity goals (Table 1). Some programs included individually tailored plans for diet and physical activity.

Table 2 summarizes the patient characteristics (with details in Supplemental Table 6). Thirty (57%) studies were restricted to participants with prediabetes, of which 21 studies used standard diagnostic criteria; 12 (23%) studies included only participants at increased risk of diabetes based on a risk score. More than three-quarters of studies included mostly overweight or obese participants and most study participants were female and at least middle-aged. Two studies were conducted in adolescents at increased risk of type 2 diabetes. These two studies are analyzed separately. None of the studies reported any long-term harms directly related to the D&PA programs.

Incident Diabetes

Sixteen studies that compared D&PA to usual care reported new-onset diabetes (6–9;22–33); two studies compared two programs each to usual care. All but three were randomized trials (9;22;26). Incident diabetes was reported between 1 and 23 years from the start of the programs (Figure 1). Across studies, between 0% (at 1 year) and 73% (at 23 years) of participants in the D&PA programs developed diabetes. At all time points participants in the D&PA programs were less likely to develop diabetes. Across all studies, the summary RR for incident diabetes was 0.59 (95% CI 0.51, 0.66) with no statistical heterogeneity. The median risk difference across studies was –11 percentage points (IQI –16, –5). Funnel plot analysis did not find different effects between larger and smaller studies (Harbord test $P=0.27$).

Both the U.S. Diabetes Prevention Program (DPP) (6) and the Finnish Diabetes Prevention Study (DPS) (7) found statistically larger effects in older participants; but while DPS found a nonsignificant effect in the youngest age group (<51 years), DPP found statistically significant effects in all age groups. Neither study found differences by sex. The DPP found no difference by race or ethnicity and DPS found no difference by educational attainment. The Japan DPP study reported a significant effect of D&PA among participants with baseline HbA1c $\geq 5.7\%$ in contrast with participants with lower baseline HbA1c, but they did not provide a statistical analysis of the difference between subgroups (34).

Comparing across studies, no significant differences were found by setting; number of sessions; program duration; whether the D&PA program was based on the DPP or DPS approach; inclusion of a weight loss goal, individual or group diet or exercise sessions (analyzed separately); or individually tailored *diet* plans, diet or exercise counselors (analyzed separately). The 11 programs that included an individually tailored *exercise* plan (RR 0.53; 95% CI 0.45, 0.63) had a possibly greater effect than the five that did not (RR 0.67; 95% CI 0.55, 0.81; P for interaction=0.070).

Six studies directly compared more versus less intensive D&PA programs (28;45;47;48;50;56). Compared with less intensive programs, more intensive programs had more sessions (four studies); weight loss, diet, or exercise goals (three studies), or—in one study each—a maintenance phase, more intensive diet and exercise plans, an exercise physiologist, individual contact sessions, or in-person (vs. DVD) sessions. All five studies that reported at least one case of incident diabetes found lower diabetes incidence with more intensive program (RR 0.28 to 0.56), but in only one study was this statistically significant (50) (Supplemental Figure 2).

Reversion to Normoglycemia

Six studies (five trials, one non-randomized study) that compared D&PA to usual care reported reversion to normoglycemia as early as 1 year from the start of the intervention (Figure 2) (6;22–25;32). Across studies, between 20% (at 2 years) and 52% (at 6 years) of participants in the D&PA programs reverted to normoglycemia. At 3 years (four studies) and across time points, the summary RR for achieving normoglycemia were statistically significant, with an overall summary RR of 1.53 (95% CI 1.26, 1.71), with no statistical heterogeneity. The median risk difference across studies was 12 percentage points (IQI 6, 14). No within-study subgroup differences were reported and no between-study subgroup differences were found. Three studies directly compared more versus less intensive programs (45;47;48), all of which favored more intensive programs (RR 1.58 to 2.11), two of which were statistically significant (47;48) (Supplemental Figure 3).

Clinical events

Three long-term studies reported all-cause mortality, two of which also reported cardiovascular mortality with no consistent pattern of results. The Da Qing study reported lower risk for all-cause mortality (hazard ratio [HR] 0.71; 95% CI 0.51, 0.99) with D&PA after 23 years (10), but this effect was restricted to women and was not significant at earlier time points (HR 1.33 at 6 years and 0.96 at 20 years) (8). Knowler et al. (DPP) found no effect at 3 years (RD –0.6/1000 person-years) and Tuomilehto et al. (DPS) found no effect at 10 years (HR 0.57; 95% CI 0.21, 1.58) (6;105). Regarding cardiovascular death, neither Da Qing (HR 0.83; 95% CI 0.48, 1.40) at 20 years nor DPP (RR 0.50; 95% CI 0.09, 2.73) at 3 years found significant effects (10; 105). The Da Qing study reported a reduction in severe retinopathy (HR 0.53; 95% CI 0.29, 0.99) (71). Limited evidence found no significance for other clinical outcomes (cardiovascular events [2 studies], nephropathy, neuropathy, and retinopathy [1 study each]), often due to lack of power.

Body weight and glycemia

The 24 studies that compared D&PA programs to usual care and reported weight change all found net weight loss with D&PA programs (6;7;9;22–24;27–33;35–41;52–55), ranging from –0.2% to –10.5% of initial body weight (summary net change –2.2%; 95% CI –2.9, –1.4); however, the studies were highly statistically heterogeneous ($I^2 = 89\%$, $P_{\text{Heterogeneity}} < 0.001$) (Figure 3). Funnel plot analysis did not find different effects between larger and smaller studies (Egger test $P = 0.51$). By meta-regression, we tested the same covariables examined for *Incident Diabetes* and the only variable for which there were different effects

across studies was for program based on DPP or DPS. The 12 programs based on DPP or DPS yielded a net change of -3.0% (95% CI $-4.1, -1.9$) compared with the 13 other programs (net change -1.6% ; 95% CI $-2.5, -0.6$; $P = 0.051$ for interaction). However, heterogeneity across studies remained high (residual $I^2=95\%$). Across all 42 D&PA programs (not compared with usual care) (6;7;22;23;27–33;35–51;54–58), none of the factors explored by meta-regression yielded statistically significant differences across studies. In contrast to the across-studies analysis, six of the nine studies that directly compared more versus less intensive D&PA programs found statistically significant greater weight loss with the more intensive programs (28;44;45;47–50;56;58) (Supplemental Figure 4).

Eighteen studies that compared D&PA programs to usual care reported glycemia outcomes (6–9;23;28–32;35–40;52;53). Overall, D&PA programs improved measures of glycemia. Across studies, at follow-up durations closest to 1 year, FG improved with a summary net change of -0.12 mmol/L (95% CI $-0.20, -0.05$; 17 studies; $I^2 = 77\%$) (-2.2 mg/dL; 95% CI $-3.6, -0.9$), 2hG improved by -0.48 mmol/L (95% CI $-0.86, -0.17$; 11 studies; $I^2 = 87\%$) (-8.6 mg/dL (95% CI $-15.5, -3.1$), and HbA1c improved by -0.08% (95% CI $-0.12, -0.04$; 8 studies; $I^2 = 0\%$) (Supplemental Table 7). Funnel plot analysis found no significant small study effect with FG (Egger test $P=0.54$), but smaller studies were more likely to have large net reductions in 2hG ($P=0.003$); however, studies reporting significant effects on FG were no more likely to report 2hG results than those with nonsignificant FG effects ($P=0.21$). Across eight studies that compared more versus less intensive D&PA programs (28;43–45;48–50;56) (Supplemental Table 8), the median net change in FG was -0.11 mmol/L (full range $-0.20, 0.17$) (-2.0 mg/dL; full range $-3.6, 1.8$), favoring more intensive programs; however, in only one study was the difference statistically significant (56). Four studies found a median net change of -0.37 mmol/L (full range $-0.6, -0.2$) (-6.7 mg/dL; full range $-11, -3.6$) in 2hG, favoring more intensive programs (44;45;48;50); in two studies the difference was significant (48;50). None of these studies reported on HbA1c.

Across the 31 D&PA programs (not compared with usual care) in 24 studies that reported FG (6–9;23;28–32;36–39;43–46;48–50;52;53;56), there were differences based on whether individual diet sessions and diet counselors were included. Adjusting for follow-up duration, programs with individual diet sessions ($n=25/31$) or with diet counselors ($n=22/31$) yielded larger decrements in FG (individual sessions: -0.24 vs. -0.02 mmol/L [-4.3 vs. -0.4 mg/dL], $P=0.020$; counselors: -0.25 vs. -0.07 mmol/L [-4.5 vs. -1.3 mg/dL; $P=0.034$).

Blood pressure and lipids

Across 17 studies comparing D&PA programs to usual care (6;7;9;22;23;28;29;32;33;35–39;52–54), at follow-up durations closest to 1 year, D&PA improved BP (systolic: net change = -1.6 mmHg; 95% CI $-2.7, -0.5$; $I^2 = 45\%$) (diastolic: net change = -1.6 mmHg; 95% CI $-2.5, -0.8$; $I^2 = 73\%$) (Supplemental Table 9). No evidence of small study effects was found (Egger test = 0.51 systolic, 0.83 diastolic) Across 14 studies, D&PA also statistically significantly improved lipid levels (7;9;22;23;28;29;33;35–39;52;53): total cholesterol (net change = -0.05 mmol/L; 95% CI $-0.12, -0.002$ [-1.8 mg/dL (95% CI $-4.6, -0.1$); 12 studies; $I^2 = 0\%$), LDL (net change = -0.09 mmol/dL; 95% CI $-0.17, -0.01$ [-3.3

mg/dL; 95% CI -6.4, -0.3]; 8 studies; $I^2 = 0\%$), HDL (net change = 0.03 mmol/L; 95% CI 0.02, 0.05 [1.2 mg/dL; 95% CI 0.7, 1.7]; 12 studies; $I^2 = 0\%$), and triglycerides (net change = -0.07 mmol/L; 95% CI -0.14, -0.02 [-6.5 mg/dL; 95% CI -12.7, -1.8; 13 studies; $I^2 = 38\%$) (Supplemental Table 10). No evidence of small study effects was found (Egger test = 0.17 total cholesterol, 0.75 HDL, 0.12 triglycerides).

Virtual programs

Five studies evaluated programs that were conducted via web-tools, social networking, email, text messaging, video, or a combination of these, with no in-person sessions (28;33;41;42;88). One study (28) found smaller, but still significant improvements in weight and FG with a DVD compared with an in-person program: weight -5% vs. -7% from baseline; FG -2.7 versus -4.2 mg/dL. Two studies (41;42) found similar effects on weight loss as found in studies with in-person sessions: -3% to -5% from baseline. One study in India (33) found that an intervention relying on text messages was effective compared to usual care, with lesser diabetes incidence over 2 years (18% vs. 27%; HR=0.64; 95% CI 0.45, 0.92) with statistically significant net differences in HDL and triglycerides, but not weight, BP, or total cholesterol. The fifth study (88), in adolescents, however, found no effect on weight (although, this was also true for a similar program with group sessions).

Programs in Adolescents

Two studies were conducted in adolescents. In the study by Savoye et al. (102), adolescents who participated in twice-a-week group sessions were significantly more likely to revert to normoglycemia, lose weight, and have lower FG and BP compared with a control group, but there was no change in lipid profile, except triglycerides. None developed diabetes during the 6 month follow-up period. The study by Patrick et al. (88) evaluated three different programs (web, web and text message, and web and group session programs) and reported no difference in weight loss compared with a control group after 6 and 12 months. The study did not report incident diabetes or FG outcomes.

Discussion

Across the wide spectrum of D&PA programs, there is strong evidence of effectiveness in reducing new-onset diabetes. Across 16 studies, participants in D&PA programs were consistently about 40% less likely to develop diabetes, but this outcome was evaluated in only a minority (30%) of studies. D&PA programs also increase the likelihood of reverting to normoglycemia and they improve diabetes and cardiometabolic risk factors, including overweight, high blood glucose, high blood pressure, and abnormal lipid profile. The effectiveness of these programs on cardiovascular disease, diabetes-related complications, and death is yet to be determined since few studies reported these outcomes.

During protocol development, we searched Medline and the Cochrane Database of Systematic Reviews for pertinent systematic reviews; none was found that was sufficiently up-to-date and that evaluated the breadth of outcomes and range of analyses evaluated in the current review. The most comprehensive review was a health technology assessment by Gillett et al. whose search was conducted in 2011 (121), but also included diet or exercise

interventions (not in combination); nine randomized trials were included. An updated search found three similar, but more restrictive reviews published since 2013. They focused on narrower subsets of studies in adults. Schellenberg et al. included nine randomized trial of D&PA programs that had at least one other component (122). Dunkley et al. included 25 studies (11 randomized trials) of D&PA programs that explicitly translated previous efficacy trials into the community settings (123), but included studies of a broader population (e.g., obese or sedentary). Aguiar et al. included only eight studies (5 randomized trials) of D&PA interventions that included both aerobic and resistance training (124). Both reviews found similar effects on weight loss, and the latter also on FG. In meta-regression, Dunkley et al. found larger changes in weight with better alignment with lifestyle intervention attributes (123).

The evidence suggests that higher intensity programs lead to greater weight loss and reduction in new-onset diabetes. While evaluated programs were too different from each other to draw firm conclusions about the unique contributions of specific program components, results from 12 studies that directly compared programs showed that people who received more intensive programs (based on features such as number of sessions, individual sessions, and additional personnel) lost more weight and were less likely to develop diabetes. The studies that compared programs with controls had very similar effects on diabetes risk across studies; therefore, no differences based on differences in their programs could be ascertained. However, across all studies, programs that provided individual (vs. group) diet sessions resulted in greater reductions in FG, as did programs that used diet counselors (vs. no diet counselors). Programs based on DPP or DPS (which were more intensive than many other programs) resulted in greater weight loss. More information on virtual delivery will be useful to increase the reach of effective D&PA programs.

Based on evidence from two of the larger studies (the U.S. DPP and the Finnish DPS), findings appear to be applicable to wide populations (in Western countries) across race and ethnicity, socioeconomic status, risk factor status, and other demographic features. With the exception of programs in two studies, all programs were applied in adults; therefore, our results may not apply in children and adolescents. However, it is likely that the benefit of D&PA programs is applicable to younger individuals at risk for type 2 diabetes because the disease mechanisms are shared between adults and children. Although most cases of diabetes in children are due to type 1 diabetes, nearly all cases of diabetes that develop from pre-diabetes (being at increased risk of diabetes) are due to type 2 diabetes. Key aspects of the pathophysiology of type 2 diabetes are similar in individuals of all ages; thus, the programs are likely to be effective regardless of age, assuming the programs are effective at changing children's diet and physical activity. The one in-person program conducted in adolescents was similarly effective as programs conducted in adults; however, the other study in adolescents of various virtual programs found no effect on weight.

Additional studies comparing D&PA programs to usual care (no program), will likely not change the overall conclusion about the D&PA programs' effectiveness with the exception of programs for children and adolescents and, possibly, in specific populations or settings where there are gaps in data. However, there are several areas that would benefit from future research in this area. Because the available programs were highly heterogeneous with many

features included, all of which likely interacted with each other, we were unable to explain the observed heterogeneity by whether programs included specific features. Furthermore, despite often protracted descriptions of the interventions, articles often failed to clearly identify who led the interventions, what the goals were, or to provide other details to one to reproduce the intervention. Future studies that compare specific program features are needed to clarify which features (eg, individual versus group sessions, few versus many sessions, differently trained counselors) optimize the effectiveness of the programs and which are less critical. It is also unclear what is the most effective way to structure the maintenance phase to help program participants maintain their improvements. Additionally, with the proliferation of mobile devices and applications, the effectiveness of virtual programs needs to be further investigated. Importantly, long-term follow-up of (existing) community-based programs are needed to evaluate the durability of the programs' effects and their effects on clinical outcomes. Although this review did not specifically address participant attrition, a better understanding is needed of what typical attrition rates are, to understand the reasons program participants drop out, and to develop methods to retain them in the programs.

In conclusion, combined D&PA promotion programs are effective for people at increased risk of type 2 diabetes for reducing new-onset diabetes, increasing reversion to normoglycemia and improving diabetes and cardiometabolic risk factors. Programs are effective across a wide range of program features but more intensive interventions appear to be more effective. Further research is needed to discern which specific program features are most important.

Supplementary Material

Refer to Web version on PubMed Central for supplementary material.

Acknowledgements1

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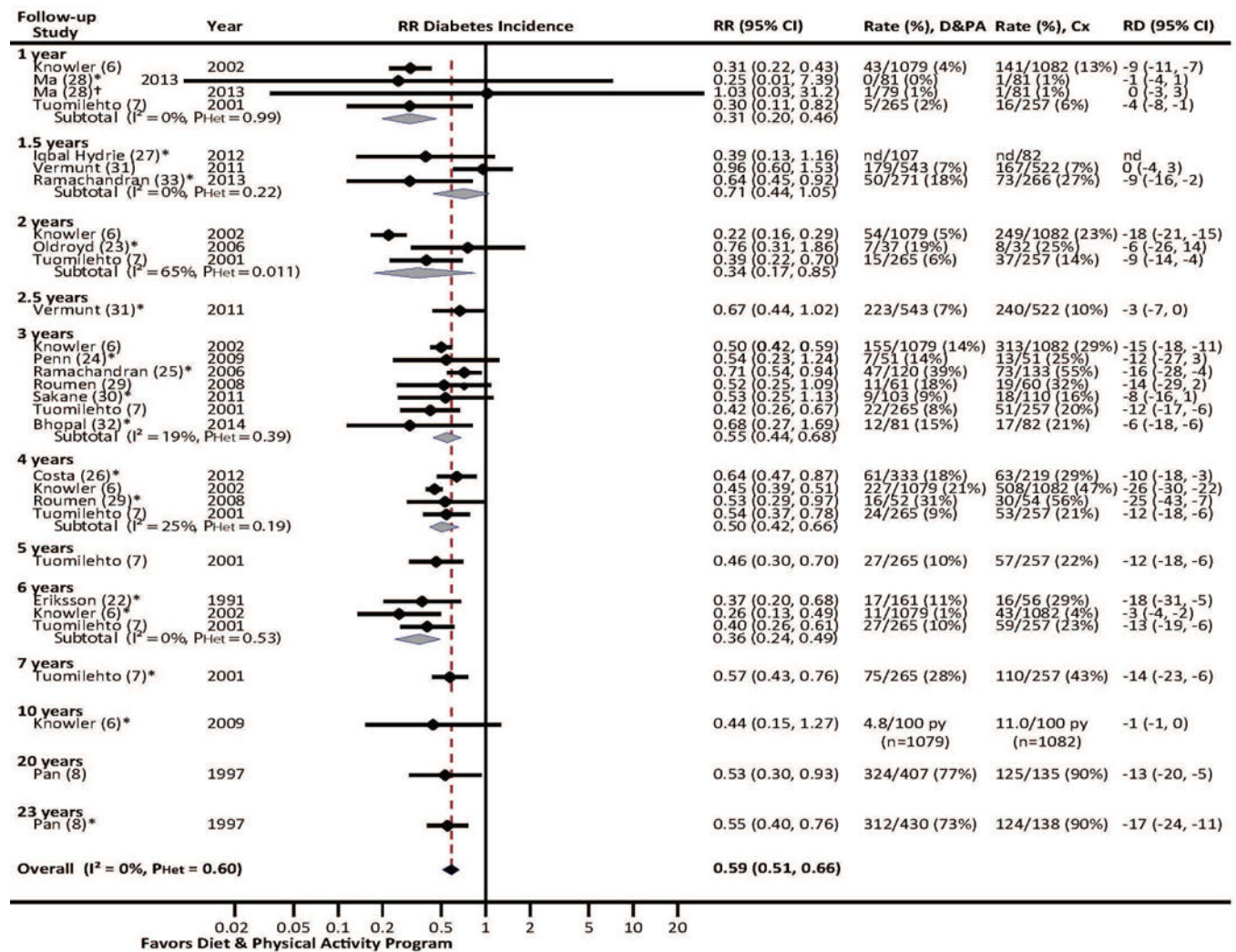


Figure 1.

Random-effects model meta-analysis of RR of incident diabetes in at-risk participants in combined diet and physical activity promotion programs vs. usual care.

For the meta-analysis of the overall summary risk ratio (RR) of incident diabetes (black diamond), the longest duration of follow-up from each study was used, as indicated by the asterisks and the grey boxes which correspond to the weight of each study in the overall meta-analysis. Subgroup meta-analyses by follow-up time (open diamonds) were conducted for time points with data from at least three studies. CI = confidence interval, Cx = control (usual care), D&PA = combined diet and physical activity promotion program, P_{Het} = Chi-squared P value of heterogeneity, RD = risk difference, RR = risk ratio.

* Included in overall meta-analysis.

† To avoid biased meta-analyses due to including correlated analyses, this comparison between the lower intensity intervention and control was excluded from meta-analysis.

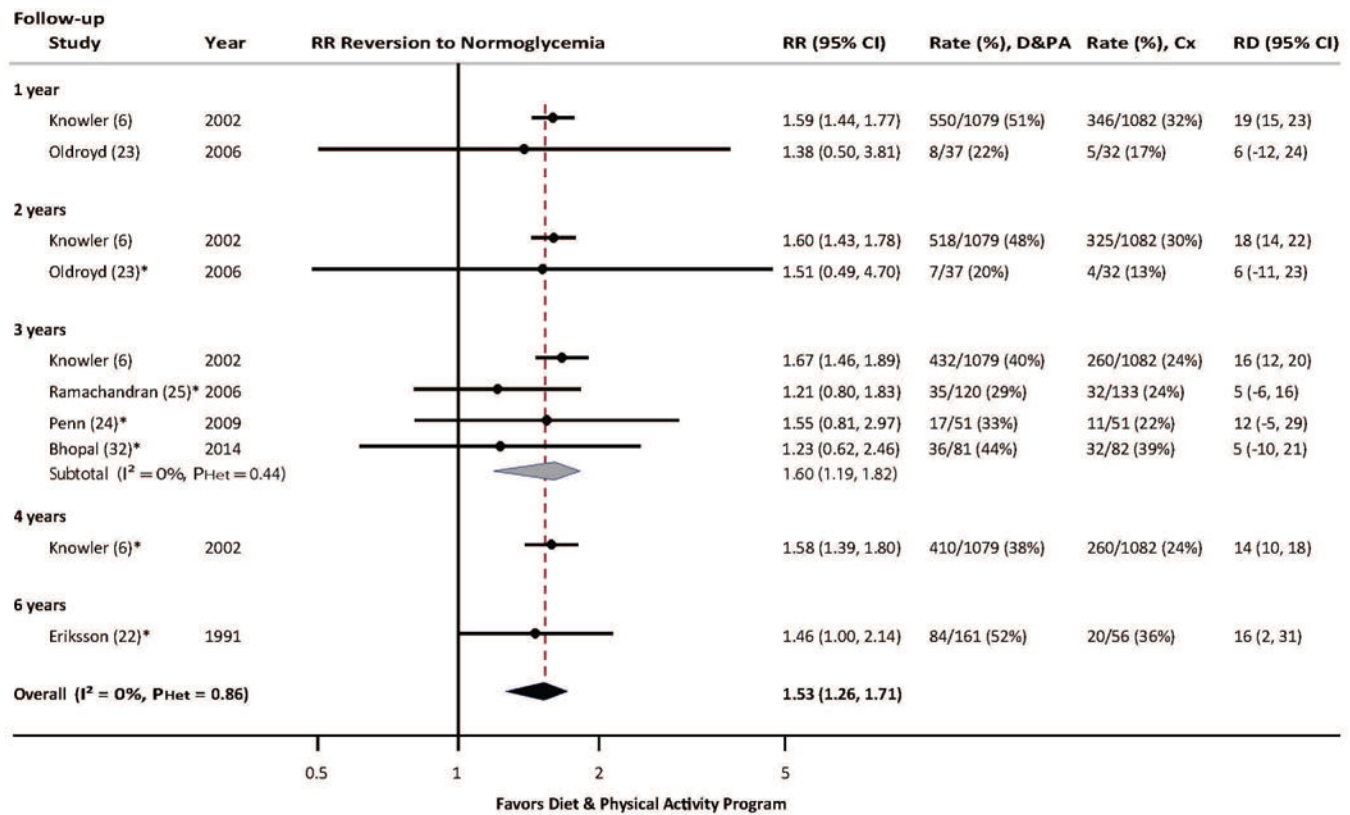
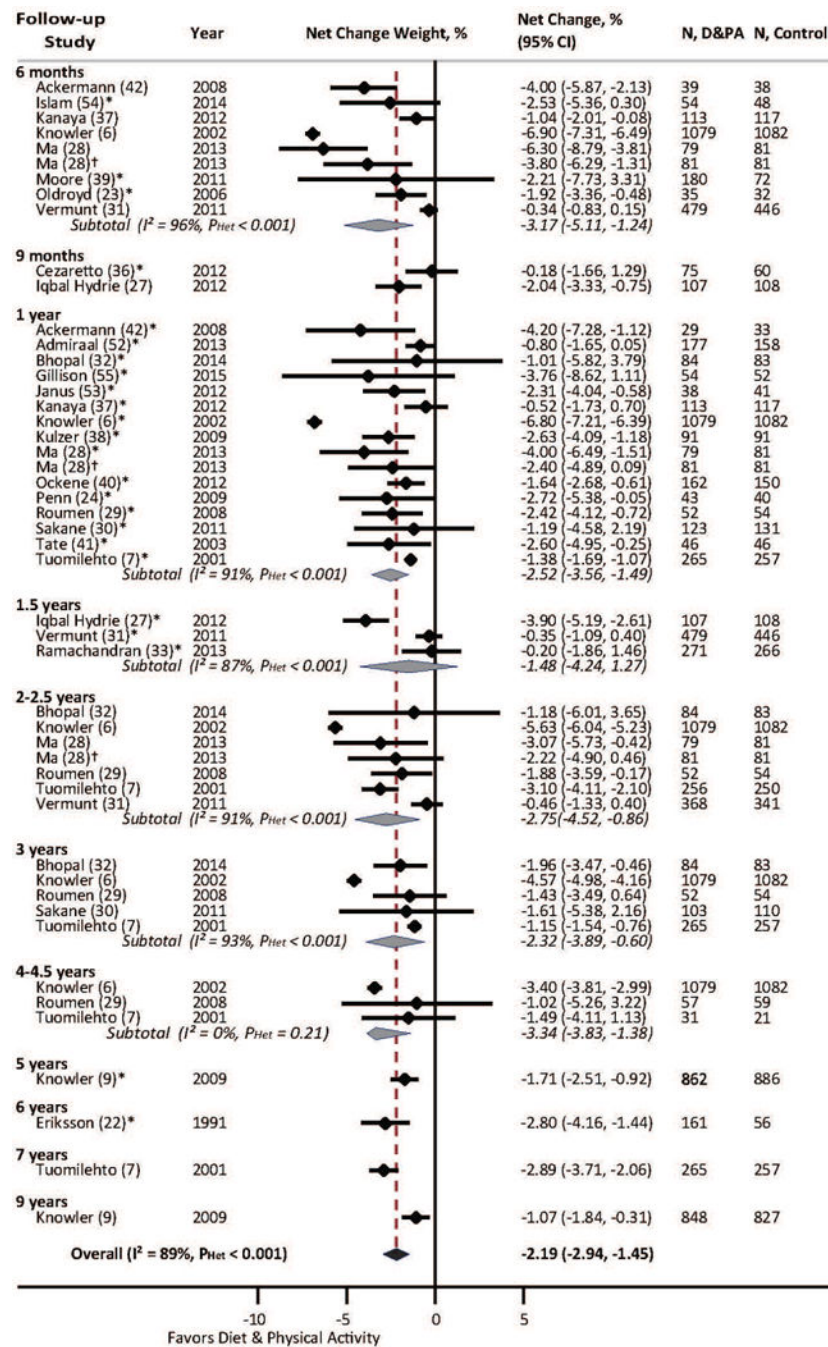


Figure 2.
Random-effects model meta-analysis of RR of reversion to normoglycemia in at-risk participants in combined diet and physical activity promotion programs vs. usual care. See Figure 1 legend.

* Included in overall meta-analysis.

**Figure 3.**

Random-effects model meta-analysis of net percent change in weight (from baseline) in at-risk participants in combined diet and physical activity promotion programs vs. usual care. See Figure 1 legend. Study data closest to 1 year follow-up were included in the overall meta-analysis.

* Included in overall meta-analysis.

† To avoid biased meta-analyses due to including correlated analyses, this comparison between the lower intensity intervention and control was excluded from meta-analysis.

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Table 1

Characteristics of Combined Diet and Physical Activity Promotion Programs

Category, No. Programs/Studies	Characteristic	Median (IQI) [Range] or No. of Programs (%)
No. sessions, 67 programs	Core	10 (6–16) [0*–72]
	Maintenance, 28 programs	6 (1.5–12) [0*–24]
	Total	15 (6.5–24.5) [0*–72]
Program duration, 67 programs	Core	6 mo (5–12) [1–60]
	Maintenance, 28 programs	12 (7–18) [4–68]
	Total	12 mo (10–27) [3–72]
Program design [†] , 67 programs	Nominally based on DPP or DPS	27 (40%)
Weight loss goal [†] , 67 programs		42 (63%)
Diet intervention [†] , 67 programs	Individual sessions	40 (60%)
	Group sessions	41 (61%)
	Individual and group	24 (36%)
	Individually tailored diet plan	16 (24%)
	Diet goal	19 (28%)
	Diet counselor	29 (43%)
	Physical activity intervention [†] , 67 programs	41 (61%)
Physical activity intervention [†] , 67 programs	Group sessions	39 (58%)
	Individual and group	24 (36%)
	Individually tailored exercise plan	23 (34%)
	Exercise goal	32 (48%)
	Exercise counselor	18 (27%)
	Counselors [†] , 51 programs	37 (73%)
	Exercise therapist	26 (51%)
Counselors [†] , 51 programs	Nurse	15 (29%)
	Layperson	13 (25%)
	Physician	8 (16%)
	Diabetes educator	3 (6%)
	Country, 53 studies	22 (42%)
	Western Europe/Australia	22 (42%)
	Japan	3 (6%)
Country, 53 studies	Middle income [‡]	6 (11%)
	Setting, 41 studies	12 (29%)
	Healthcare system	25 (61%)
	Worksite	0
	Multiple	4 (10%)
Location, 53 studies	Urban	25 (47%)
	Regional	21 (40%)
	Suburban	2 (4%)
	Rural	1 (2%)

Category, No. Programs/Studies	Characteristic	Median (IQI) [Range] or No. of Programs (%)
	Mixed	4 (8%)

DPP = U.S. Diabetes Prevention Program Trial, DPS = Finnish Diabetes Prevention Study, IQI = interquartile interval, mo = months.

* In some programs, the contacts were by telephone, email, internet, or video only.

[†] Likely underestimates due to inadequate or unclear reporting in articles.

[‡] India 3, Brazil 1, China 1, Pakistan 1.

Table 2

Characteristics of Study Participants

Category, No. Studies	Characteristic	Median (IQI) [Range] or No. of Studies (%)
Eligibility criteria, 53 studies	Prediabetes, IGT, or IFG	30 (57%)
	By ADA/WHO criteria	21 (40%)
	At increased risk for DM (by risk score)	12 (23%)
	Prediabetes or at increased risk for DM	4 (8%)
	Prediabetes or at increased risk for CVD	4 (8%)
	Metabolic syndrome \pm prediabetes	3 (6%)
Body weight, 47 studies	Mean BMI, kg/m ²	31.2 (28.1, 33.6) [23.8, 39.7]
Hypertension, 4 studies	% participants	34.5 [30.6, 50]
Female, 39 studies	% participants	65.3 (50.3, 73.9) [13.5, 90.5]
Age, 39 studies [†]	Mean years	53.6 (48, 57) [43.1, 65.0]
Ethnicity [‡]		
13 studies	White, % participants	74 [18, 89]
10 studies	Black/African-American, % participants	18 [12, 39]
8 studies	Hispanic/Latino, % participants	13 [3, 38]
5 studies	East Asian, % participants	100
6 studies	Southeast Asian, % participants	100
4 studies	Asian/Pacific Islander, % participants	4, 5, 15, 17 [§]
4 studies	Native American, % participants	1, 3, 6, 100 [§]
Education		
9 studies	<High school (or equivalent), % participants	14 (11, 33) [5, 64]
20 studies	High school (or some college), % participants	30 (21, 48) [10, 69]
11 studies	Bachelors degree (or equivalent), % participants	28 (20, 37) [14, 52]
4 studies	Graduate degree (or equivalent), % participants	13, 15, 16, 35 [§]

DPP = U.S. Diabetes Prevention Program Trial, DPS = Finnish Diabetes Prevention Study, IQI = interquartile interval, mo = months.

* In some programs, the maintenance period contacts were by telephone or email only.

[†] Excluding 2 studies in adolescents.

[‡] Excluding counts of studies with 0% of an ethnicity.

[§] List of percentages among relevant studies.